PATENT COOPERATION TREATY

PCT

INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY

(Chapter II of the Patent Cooperation Treaty)

(PCT Article 36 and Rule 70)

1	cant's or agent's file reference	FOR FURTHER ACTIO	ON S	See Form PCT/IPEA/416	
B-15300 PCT International application No.		International filing date (day)	month/year)	Priority date (day/month/year) .	
1		25.02.2005		26.02.2004	
ł	national Patent Classification (IPC) or r BB37/00	lational classification and IPC			
Applicant PHARMA MEDICAL LIMÍTED et al					
1.	 This report is the international preliminary examination report, established by this International Preliminary Examining Authority under Article 35 and transmitted to the applicant according to Article 36. 				
2.	This REPORT consists of a total of 5 sheets, including this cover sheet.				
3.					
 a. sent to the applicant and to the International Bureau) a total of sheets, as follows: sheets of the description, claims and/or drawings which have been amended and are the basis of thi and/or sheets containing rectifications authorized by this Authority (see Rule 70.16 and Section 607 Administrative Instructions). 				s follows:	
				e fulle 70.10 and Section Sov St. and	
	sheets which supersede earlier sheets, but which this Authority considers contain an amendment that goes beyond the disclosure in the international application as filed, as indicated in item 4 of Box No. I and the Supplemental Box.				
	b. (sent to the International Bureau only) a total of (indicate type and number of electronic carrier(s)), containing a sequence listing and/or tables related thereto, in computer readable form only, as indicated in the Supplemental Box Relating to Sequence Listing (see Section 802 of the Administrative Instructions).				
4.	This report contains indications	relating to the following item	s:		
"	_				
	<u> </u>	JINOT			
	☐ Box No. II Priority ☐ Box No. III Non-establish	ment of oninion with regard t	to novelty, inventive	step and industrial applicability	
1			••		
	 ☐ Box No. IV Lack of unity of invention ☑ Box No. V Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement 				
	☐ Box No. VI Certain docum				
	☐ Box No. VII Certain defec	s in the international applica	ation		
	Box No. VIII Certain observations on the international application				
	e of submission of the demand		Date of completion of the	nis report	
Dat	e or samillasion of the delitatio] -	· •	•	
05.	.09.2005	1	18.01.2006		
Name and mailing address of the international preliminary examining authority:		onal	Authorized Officer		
-	European Patent Office D-80298 Munich Tel. +49 89 2399 - 0 Tx: 52	3656 epmu d	Gerber, M	· 11.00 (1.0	
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INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY

International application No. PCT/IB2005/000506

	Box No. I Basis of the report				
_ 1		s report is based on the international application in the language in which it was			
1,	With regard to the language , this report is based on the international application in the language in which it was filed, unless otherwise indicated under this item.				
	☐ This report is based on trans	This report is based on translations from the original language into the following language, which is the language of a translation furnished for the purposes of:			
		☐ international search (under Rules 12.3 and 23.1(b))			
	publication of the internat	ional application (under Rule 12.4)			
	☐ international preliminary e	examination (under Rules 55.2 and/or 55.3)			
2.	. With regard to the elements* of t have been furnished to the receiv report as "originally filed" and are	the international application, this report is based on (replacement sheets which ving Office in response to an invitation under Article 14 are referred to in this e not annexed to this report):			
	Description, Pages				
	1-18	as originally filed			
	Claims, Numbers				
	1-36	as originally filed			
	Drawings, Sheets				
	1/7-7/7	as originally filed			
	☐ a sequence listing and/or any	y related table(s) - see Supplemental Box Relating to Sequence Listing			
3.	. The amendments have resul	ited in the cancellation of:			
	the description, pages				
	☐ the claims, Nos.☐ the drawings, sheets/figs				
	□ the sequence listing (specified)				
	☐ any table(s) related to sec	quence listing (specify):			
4.		shed as if (some of) the amendments annexed to this report and listed below ave been considered to go beyond the disclosure as filed, as indicated in the			
	☐ the description, pages				
	☐ the claims, Nos.☐ the drawings, sheets/figs				
	☐ the sequence listing (spec				
	any table(s) related to sec	quence listing (specify):			
	* If item 4 applies, son	me or all of these sheets may be marked "superseded."			

INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY

International application No. PCT/IB2005/000506

Box No. V Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

1. Statement

Novelty (N) Yes: Claims 1-36

No: Claims

Inventive step (IS) Yes: Claims 1-36

No: Claims

Industrial applicability (IA) Yes: Claims 1-21

No: Claims 22-36

2. Citations and explanations (Rule 70.7):

see separate sheet

10/590625 IAP9 Rec'd PCT/PTO 24 AUG 2006

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PCT/IB2005/000506

Re Item V

Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

Reference is made to the following documents:

D1: CARNEY S L: "EFFECT OF DIACETYL RHEIN ON THE DEVELOPMENT OF EXPERIMENTAL OSTEOARTHRITIS A BIOCHEMICAL INVESTIGATION" December 1996, OSTEOARTHRITIS AND CARTILAGE, BAILLIERE TINDALL, LONDON, GB, PAGE(S) 251-261, XP001056066 ISSN: 1063-4584

D2: BARBUCCI R ET AL: "Hyaluronic acid hydrogel in the treatment of osteoarthritis" December 2002, BIOMATERIALS, ELSEVIER SCIENCE PUBLISHERS BV., BARKING, GB, PAGE(S) 4503-4513, XP004377521 ISSN: 0142-9612

1. Novelty

D1 is concerned with the use of rhein in the treatment of joint disorders and D2 deals with the implementation of hyaluronic acid in the treatment of osteoarthritis.

None of these documents discloses the esterification of hyaluronic acid with rhein.

The subject-matter of claims 1-36 is novel over the cited state of the art (Article 33(2) PCT).

2. Inventive step

The hyaluronic acid esterified with rhein exhibits an improved pharmacological activity than hyaluronic acid or rhein alone (see page 17, line 14 - page 18, line 2, as well as figures 4 and 5).

The technical problem underlying the present application is the provision of a compound based on hyaluronic acid with improved pharmacological activity.

None of the cited documents cited alone or in combination would have prompted the

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skilled person seeking to solve the above-mentioned technical problem to esterify hyaluronic acid with rhein.

The subject-matter of claims 1-36 is therefore considered inventive (Article 33(3) PCT).

3. Industrial applicability

- **3.1.** The subject-matter of present claims 1-21 appears to comply with the requirements of industrial applicability as stipulated in Article 33(4) PCT.
- **3.2.** Claims 22-36 relate to subject-matter considered by this Authority to be covered by the provisions of Rule 67.1(iv) PCT. Consequently, no opinion will be formulated with respect to the industrial applicability of the subject-matter of these claims (Article 34(4)(a)(I) PCT).